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Augmentation of antiproliferative activity of interferon alfa against human bladder tumor cell lines by encapsulation of interferon alfa within liposomes.

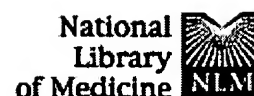
Killion JJ, Fan D, Bucana CD, Frangos DN, Price JE, Fidler IJ.

Department of Cell Biology, University of Texas M.D. Anderson Cancer Center, Houston 77030.

Present therapy for human bladder cancer includes the intravesical administration of antiproliferative agents, such as recombinant human interferon alfa (IFN- α). The administration of cytotoxic molecules encapsulated in liposomes could provide a more efficient method for such therapy. Therefore, we determined whether encapsulation of the recombinant human IFN- α hybrid BBDD within liposomes will produce antitumor effects against the human bladder cancer cell line 253J superior to those observed with free IFN- α . Adherent cells were cultured in medium alone, in medium containing different concentrations of IFN- α , or in medium containing multilamellar liposomes (phosphatidylcholine-phosphatidylserine at a molar ratio of 7:3) that encapsulated saline or IFN- α . Cell growth was determined 96-120 hours later. Additional control groups consisted of target cells cultured with free IFN- α or with IFN- α plus liposomes containing saline. Cytostasis mediated by free IFN- α alone or IFN- α in the presence of liposome-saline was identical and ranged from 0%-30% (10 IU/mL) to 45%-70% (1,000 IU/mL). Liposomes containing saline produced no effects. Liposome-encapsulated IFN- α produced significantly greater growth inhibition than free IFN- α : 40%-70% (10 IU/mL) and 80%-90% (1,000 IU/mL), respectively. Moreover, a 253J variant subline selected for resistance to free IFN- α was sensitive to IFN- α presented in liposomes. These data suggest that the encapsulation of antiproliferative agents such as IFN- α in liposomes can improve therapeutic results.

PMID: 2778824 [PubMed - indexed for MEDLINE]

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